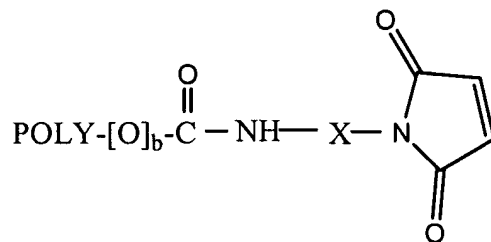


Amendments to the Claims:

The following listing reflects amendments to the claims and replaces all prior versions and listings of claims in this application.

1. (Withdrawn). A water-soluble polymer having the structure:



II

wherein:

POLY is a water-soluble polymer segment,

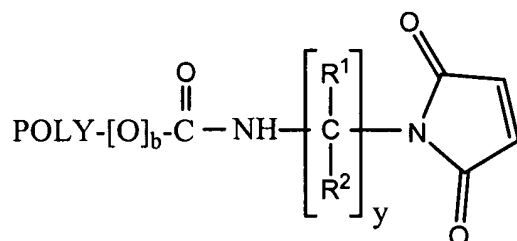
b is 0 or 1,

X is a hydrolytically stable linker comprising at least 3 contiguous saturated carbon atoms, and

said polymer is absent aromatic groups and ester linkages.

2. (Withdrawn) The polymer of claim 1, wherein X is a saturated acyclic, cyclic or alicyclic hydrocarbon chain having a total of about 3 to about 20 carbon atoms.
3. (Withdrawn) The polymer of claim 2, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, and 20.
4. (Withdrawn) The polymer of claim 3, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of: from about 3 to about 20, from about 4 to about 12, from about 4 to about 10, and from about 5 to about 8 atoms.

5. (Withdrawn) The polymer of any one of claims 2 to 4, wherein X is a linear saturated acyclic hydrocarbon chain.
6. (Withdrawn) The polymer of claim 5, wherein X is a branched saturated acyclic hydrocarbon chain.
7. (Withdrawn) The polymer of claim 6, wherein X is branched at the carbon  $\alpha$  to the maleimidyl group.
8. (Withdrawn) The polymer of claim 6, wherein X is branched at the carbon  $\beta$  to the maleimidyl group.
9. (Withdrawn) The polymer of claim 6, wherein X is branched at the carbon  $\gamma$  to the maleimidyl group.
10. (Withdrawn) The polymer of claim 5, having the structure:



III

wherein:

y is an integer from 1 to about 20;

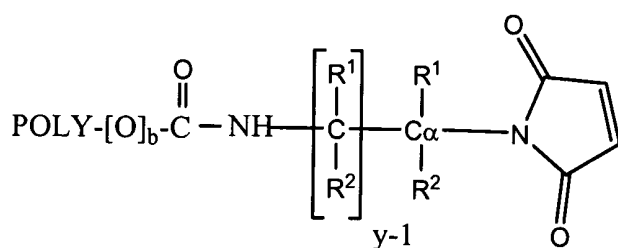
R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and

R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

11. (Withdrawn) The polymer of claim 10, wherein  $R^1$  and  $R^2$  in each occurrence is independently H or an organic radical selected from the group consisting of lower alkyl and lower cycloalkyl.

12. (Withdrawn) The polymer of claim 10, wherein  $R^1$  and  $R^2$  are both H, and Y is selected from the group consisting of 3, 4, 5, 6, 7, 8, 9, and 10.

13. (Withdrawn) The polymer of claim 10 having the structure:



### III-A

wherein at least one of  $R^1$  or  $R^2$  on  $C_\alpha$  is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and y is at least one.

14. (Withdrawn) The polymer of claim 13, wherein each of  $R^1$  and  $R^2$  on  $C_\alpha$  is independently selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

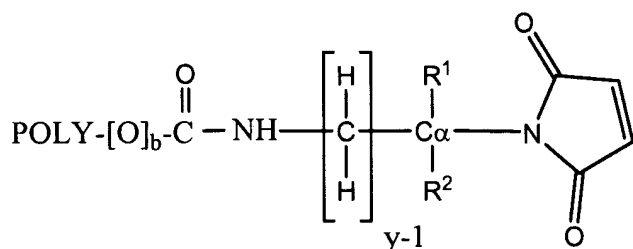
15. (Withdrawn) The polymer of claim 13, wherein all other non- $C_\alpha$   $R^1$  and  $R^2$  variables are H.

16. (Withdrawn) The polymer of claim 13 wherein at least one of  $R^1$  or  $R^2$  on  $C_\alpha$  is lower alkyl or lower cycloalkyl.

17. (Withdrawn) The polymer of claim 13; wherein  $R^2$  on  $C_\alpha$  is H.

18. (Withdrawn) The polymer of claim 17, wherein R<sup>1</sup> on C<sub>α</sub> is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, isobutyl, pentyl, cyclopentyl, hexyl, and cyclohexyl.

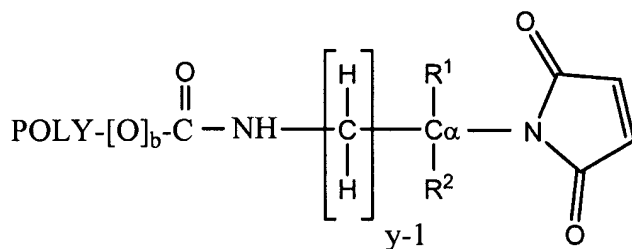
19. (Withdrawn) The polymer of claim 13 having the structure:



### III-B

wherein R<sup>1</sup> and R<sup>2</sup> is each independently alkyl or cycloalkyl.

20. (Withdrawn) The polymer of claim 13, having the structure:



### III-B

wherein R<sup>1</sup> is alkyl or cycloalkyl and R<sup>2</sup> is H.

21. (Withdrawn) The polymer of claim 19 wherein R<sup>1</sup> and R<sup>2</sup> are each independently either methyl or ethyl.

22. (Withdrawn) The polymer of claim 19, wherein R<sup>1</sup> and R<sup>2</sup> are the same.

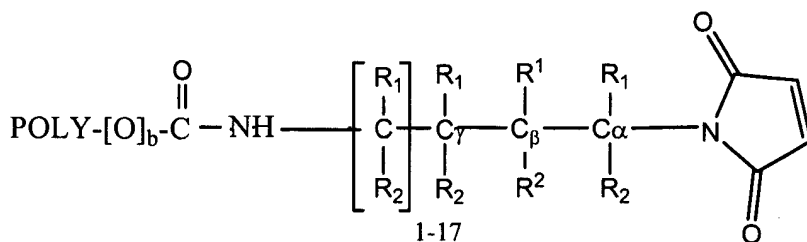
23. (Withdrawn) The polymer of claim 1, wherein said polymer possesses a hydrolytically stabilized maleimide ring.

$$\text{POLY-}[\text{O}]_b\text{-}\overset{\text{O}}{\parallel}\text{C}-\text{NH}-\left[\begin{array}{c} \text{H} \\ | \\ \text{C}_\gamma \\ | \\ \text{H} \end{array}\right]_{\gamma-2}-\begin{array}{c} \text{R}^1 \\ | \\ \text{C}_\beta \\ | \\ \text{R}^2 \end{array}-\begin{array}{c} \text{H} \\ | \\ \text{C}_\alpha \\ | \\ \text{H} \end{array}-\text{N}-\text{C}_5\text{H}_3\text{N}_2\text{O}_2$$

wherein R<sup>1</sup> and R<sup>2</sup> is each independently selected from the group consisting of H, alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl, but are not both H, and y is at least 2.

26. (Withdrawn) The polymer of claim 25, wherein R<sup>1</sup> and R<sup>2</sup> is each independently selected from the group consisting of H, methyl, ethyl, propyl, isopropyl, butyl, isobutyl, pentyl, cyclopentyl, hexyl, and cyclohexyl.

28. (Withdrawn) The polymer of claim 10, having the structure:



-6-

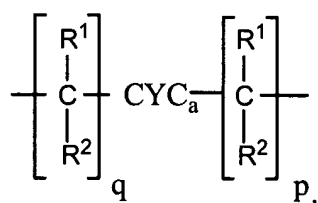
wherein at least one of  $R^1$  and  $R^2$  attached to  $C_\gamma$  is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

29. (Withdrawn) The polymer of claim 28, wherein at least one of  $R^1$  and  $R^2$  attached to  $C_\gamma$  is alkyl or cycloalkyl and all other  $R^1$  and  $R^2$  variables are H.

30. (Withdrawn) The polymer of claim 28, wherein one of the  $R^1$  variables attached to  $C_\alpha$  or  $C_\beta$  is alkyl or cycloalkyl, and all other  $R^1$  and  $R^2$  variables are H.

31. (Withdrawn) The polymer of claim 2, wherein X is a saturated cyclic or alicyclic hydrocarbon chain.

32. (Withdrawn) The polymer of claim 31, wherein X has the structure:



**IV**

and

$\text{CYC}_a$  is a cycloalkylene group having "a" ring carbons, where the value of "a" ranges from 3 to 12;

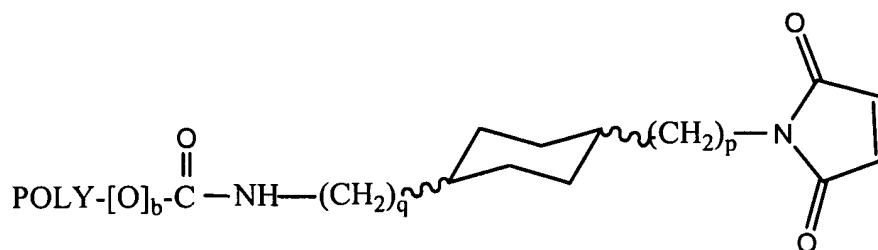
p and q are each independently 0 to 20, and  $p + q + a \leq 20$ ,

$R^1$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and

$R^2$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

33. (Withdrawn) The polymer of claim 32, wherein p and q are each independently selected from the group consisting of 0, 1, 2, 3, 4, 5, 6, 7, and 8.
34. (Withdrawn) The polymer of claim 32, wherein R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl, and R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl.
35. (Withdrawn) The polymer of claim 32, wherein a is selected from the group consisting of 5, 6, 7, 8 and 9.
36. (Withdrawn) The polymer of claim 35, wherein a is 6 and CYC<sub>a</sub> is a 1,1-, 1,2-, 1,3- or 1,4-substituted cyclohexyl ring.
37. (Withdrawn) The polymer of claim 32, wherein p and q each independently range from 0 to 4.
38. (Withdrawn) The polymer of claim 36, wherein the substituents on said substituted cyclohexyl ring are cis.
39. (Withdrawn) The polymer of claim 36, wherein the substituents on said substituted cyclohexyl ring are trans.
40. (Withdrawn) The polymer of claim 32, wherein R<sup>1</sup> and R<sup>2</sup> are H in every occurrence.

41. (Withdrawn) The polymer of claim 34, having the structure:

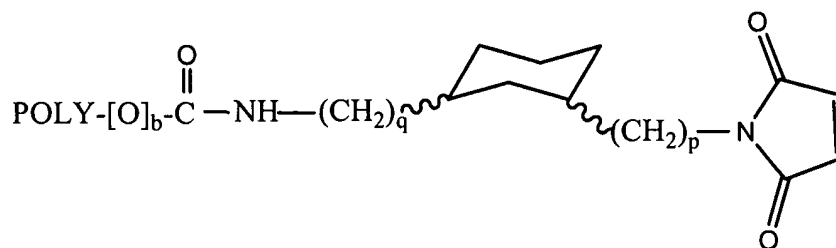


**IV-A**

wherein q and p each independently range from 0 to 6.

42. (Withdrawn) The polymer of claim 41, wherein q ranges from 0 to 6 and p is zero.

43. (Withdrawn) The polymer of claim 34, having the structure:

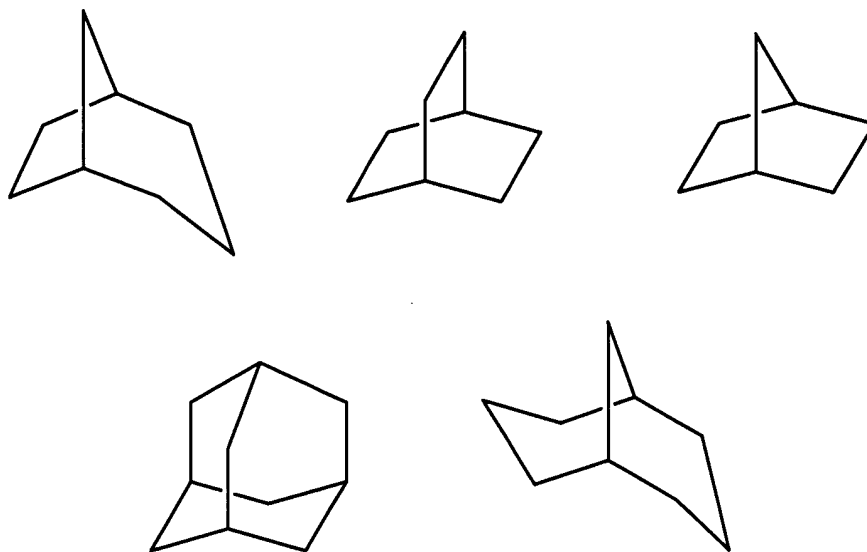


wherein q and p each independently range from 0 to 6, and the substituents on the cyclohexylene ring are either cis or trans.

44. (Withdrawn) The polymer of claim 32, wherein CYC<sub>a</sub> is bicyclic or tricyclic.

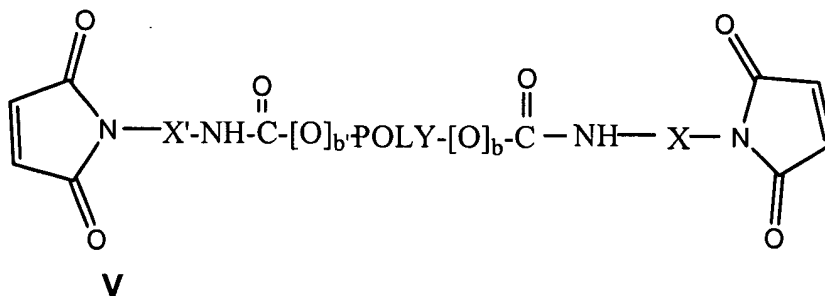


45. (Withdrawn) The polymer of claim 44, wherein  $CYC_a$  is selected from the group consisting of:



and said ring substituents are positioned at any available position on the bi or tricyclic ring.

46. (Withdrawn) The polymer of claim 1 having the structure:



wherein X and b are as previously defined,

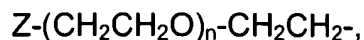
$b'$  is 0 or 1, and

$X'$  is a hydrolytically stable linker comprising at least 3 contiguous saturated carbon atoms.

47. (Withdrawn) The polymer of claim 1, wherein POLY is selected from the group consisting of a poly(alkylene oxide), poly(vinyl pyrrolidone), poly(vinyl alcohol), polyoxazoline, poly(acryloylmorpholine), and poly(oxyethylated polyol).
48. (Withdrawn) The polymer of claim 47, wherein POLY is a poly(alkylene oxide).
49. (Withdrawn) The polymer of claim 48, wherein POLY is a poly(ethylene glycol).
50. (Withdrawn) The polymer of claim 49, wherein the poly(ethylene glycol) is terminally capped with an end-capping moiety.
51. (Withdrawn) The polymer of claim 50, wherein the end-capping moiety is independently selected from the group consisting alkoxy, substituted alkoxy, alkenyloxy, substituted alkenyloxy, alkynyloxy, substituted alkynyloxy, aryloxy, and substituted aryloxy.
52. (Withdrawn) The polymer of claim 51, wherein the end-capping moiety is selected from the group consisting of methoxy, ethoxy, and benzyloxy.
53. (Withdrawn) The polymer of claim 49, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 100 daltons to about 100,000 daltons.
54. (Withdrawn) The polymer of claim 53, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 1,000 daltons to about 50,000 daltons.
55. (Withdrawn) The polymer of claim 54, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 2,000 daltons to about 30,000 daltons.
56. (Withdrawn) The polymer of claim 46, wherein said POLY is linear and the polymer is homobifunctional.

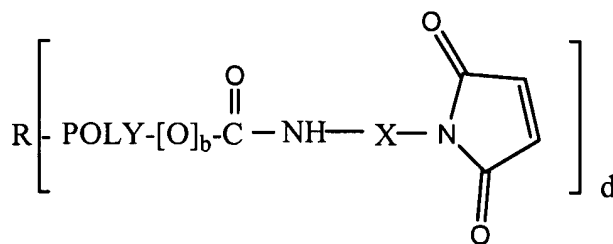
57. (Withdrawn) The polymer of claim 49, wherein said poly(ethylene glycol) has a structure selected from the group consisting of linear, branched and forked.

58. (Withdrawn) The polymer of claim 26, wherein said poly(ethylene glycol) comprises the structure:



where n is from about 10 to about 4000, and Z comprises a moiety selected from the group consisting of hydroxy, amino, ester, carbonate, aldehyde, aldehyde hydrate, acetal, ketone, ketone hydrate, ketal, alkenyl, acrylate, methacrylate, acrylamide, sulfone, thiol, carboxylic acid, isocyanate, isothiocyanate, hydrazide, urea, maleimide, vinylsulfone, dithiopyridine, vinylpyridine, iodoacetamide, alkoxy, benzyloxy, silane, lipid, phospholipid, biotin, and fluorescein.

59. (Withdrawn) The polymer of claim 1, corresponding to the structure:



**VI**

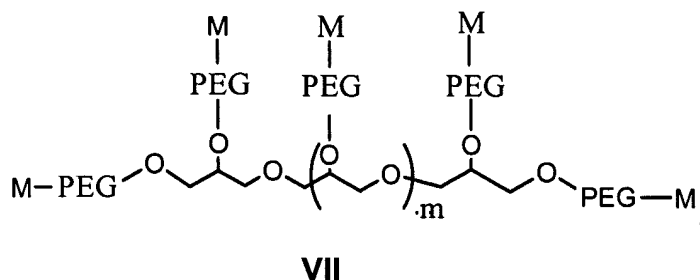
wherein:

d is an integer from 3 to about 100, and

R is a residue of a central core molecule having 3 or more hydroxyl groups, amino groups, or combinations thereof.

60. (Withdrawn) The polymer of claim 59, wherein d is an integer from 3 to about 12.

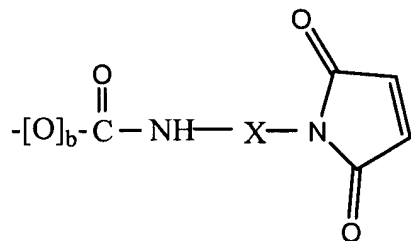
61. (Withdrawn) The polymer of claim 1, wherein POLY is a multi-arm polymer segment, and said polymer corresponds to the structure:



where

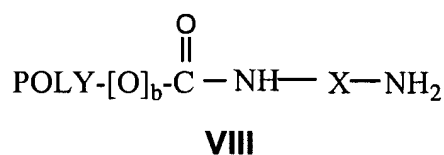
PEG is  $-(\text{CH}_2\text{CH}_2\text{O})_n\text{CH}_2\text{CH}_2-$ ,

M is :



and m is selected from the group consisting of 3, 4, 5, 6, 7, and 8.

62. (Withdrawn) A water-soluble polymer having the structure:



wherein

POLY is a water-soluble polymer segment,

b is 0 or 1,

X is a hydrolytically stable linker comprising at least 3 contiguous saturated carbon atoms,

and

said polymer is absent aromatic groups and ester linkages.

63. (Withdrawn) The polymer of claim 62, wherein X is a saturated acyclic, cyclic or alicyclic hydrocarbon chain having a total of about 3 to about 20 carbon atoms.

64. (Withdrawn) The polymer of claim 62, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, and 20.

65. (Withdrawn) The polymer of claim 64, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of: from about 3 to about 20, from about 4 to about 12, from about 4 to about 10, and from about 5 to about 8 atoms.

66. (Withdrawn) The polymer of claim 63, wherein X is a linear saturated acyclic hydrocarbon chain.

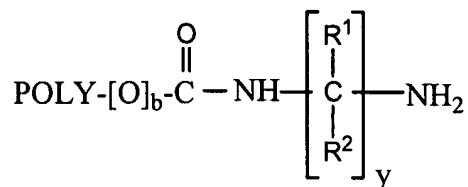
67. (Withdrawn) The polymer of claim 63, wherein X is a branched saturated acyclic hydrocarbon chain.

68. (Withdrawn) The polymer of claim 67, wherein X is branched at the carbon  $\alpha$  to the maleimidyl group.

69. (Withdrawn) The polymer of claim 67, wherein X is branched at the carbon  $\beta$  to the maleimidyl group.

70. (Withdrawn) The polymer of claim 67, wherein X is branched at the carbon  $\gamma$  to the maleimidyl group.

71. (Withdrawn) The polymer of claim 63, having the structure:



**VIII-A**

wherein:

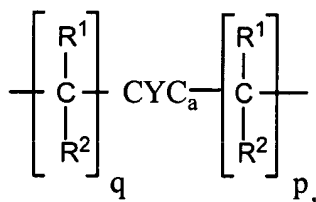
y is an integer from 1 to about 20;

R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and

R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

72. (Withdrawn) The polymer of claim 62, wherein X is a saturated cyclic or alicyclic hydrocarbon moiety.

73. (Withdrawn) The polymer of claim 72, wherein X has the structure:



**VIII-B**

and

CYC<sub>a</sub> is a cycloalkylene group having "a" ring carbons, where the value of "a" ranges from 3 to 12;

p and q are each independently 0 to 20, and p + q + a ≤ 20,

$R^1$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkenylcycloalkyl, and substituted alkenylcycloalkyl, and

$R^2$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkenylcycloalkyl, and substituted alkenylcycloalkyl.

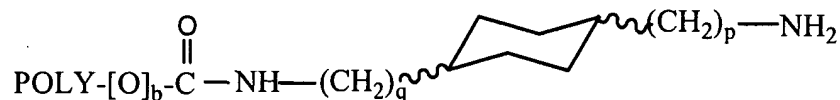
74. (Withdrawn) The polymer of claim 73, wherein p and q are each independently selected from the group consisting of 0, 1, 2, 3, 4, 5, 6, 7, and 8.

75. (Withdrawn) The polymer of claim 73, wherein  $R^1$ , in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl, and  $R^2$ , in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl.

76. (Withdrawn) The polymer of claim 73, wherein a is selected from the group consisting of 5, 6, 7, 8 and 9.

77. (Withdrawn) The polymer of claim 76, wherein a is 6 and  $CYC_a$  is a 1,1-, 1,2-, 1,3- or 1,4-substituted cyclohexylene ring.

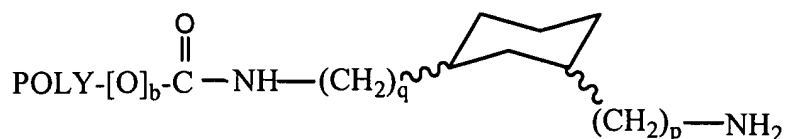
78. (Withdrawn) The polymer of claim 77, having the structure:



**VIII-C**

wherein q and p each independently range from 0 to 6, and the substituents on said cyclohexylene ring are either cis or trans.

79. (Withdrawn) The polymer of claim 77, having the structure:



### VIII-D

wherein q and p each independently range from 0 to 6, and the substituents on said cyclohexylene ring are either cis or trans.

80. (Withdrawn) The polymer of claim 73, wherein  $\text{CYC}_a$  is bicyclic or tricyclic.

81. (Withdrawn) A method for forming a maleimide terminated polymer, said method comprising the steps of:

- a. reacting  $\text{POLY}-[\text{O}]_b-\text{C}(\text{O})-\text{LG}$  **(IX)** with  $\text{NH}_2-\text{X}-\text{NH}_2$  **(XII)** under conditions effective to form  $\text{POLY}-[\text{O}]_b-\text{C}(\text{O})-\text{H}_2\text{N}-\text{X}-\text{NH}_2$  **(X)**, and
- b. converting  $\text{POLY}-[\text{O}]_b-\text{C}(\text{O})-\text{H}_2\text{N}-\text{X}-\text{NH}_2$  **(X)** into  $\text{POLY}-[\text{O}]_b-\text{C}(\text{O})-\text{HN}-\text{X}-\text{MAL}$  **(II)**,

wherein:

POLY is a water-soluble polymer segment,

b is 0 or 1,

X is a hydrolytically stable linker comprising at least 3 contiguous saturated carbon atoms,

LG is a leaving group,

MAL is maleimide,

and said maleimide terminated polymer is absent aromatic groups and ester linkages.



82. (Withdrawn) The method of claim 81, wherein said LG is selected from the group consisting of halide, N-hydroxysuccinimide, N-hydroxybenzotriazole, para-nitrophenolate.

83. (Withdrawn) The method of claim 81, wherein one of said amino groups in said  $\text{NH}_2\text{-X-NH}_2$  reagent is in protected form.

84. (Withdrawn) The method of claim 81, wherein said reacting step is carried out in an organic solvent selected from the group consisting of acetonitrile, chloroform, dichloromethane, benzene, toluene, xylene, acetone, tetrahydrofuran (THF), dimethylformamide (DMF), and dimethylsulfoxide.

85. (Withdrawn) The method of claim 81, wherein said reacting step is conducted under an inert atmosphere.

86. (Withdrawn) The method of claim 81, wherein said reacting step is conducted at a temperature in the range of about 0 to 100° C.

87. (Withdrawn) The method of claim 81, wherein said reacting step is conducted in the presence of a base selected from the group consisting of triethylamine, pyridine, 4-(dimethylamino)pyridine, and sodium carbonate.

88. (Withdrawn) The method of claim 83, further comprising after said reacting step, deprotecting the amino group in  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-NH}_2$ .

89. (Withdrawn) The method of claim 81, further comprising the step of purifying the product from step (a) prior to said converting step.

90. (Withdrawn) The method of claim 89, wherein said purifying comprises purifying the product by column chromatography.

91. (Withdrawn) The method of claim 89, wherein said purifying comprises purifying the product by ion exchange chromatography.

92. (Withdrawn) The method of claim 81, wherein said converting step comprises reacting  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-NH}_2$  with a reagent selected from the group consisting of N-methoxycarbonylmaleimide, exo-7-oxa[2.2.1]bicycloheptane-2,3-dicarboxylic anhydride, and maleic anhydride under conditions suitable for forming  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-MAL}$  in a reaction mixture.

93. (Withdrawn) The method of claim 92, wherein when said reagent is N-methoxycarbonylmaleimide, and said converting step is carried out in water or an aqueous mixture of water and a water miscible solvent.

94. (Withdrawn) The method of claim 92, wherein said reagent is maleic anhydride, and said converting step comprises reacting  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-NH}_2$  with maleic anhydride under conditions effective to form  $\text{POLY-[O]}_b\text{-C(O)-NH-X-NH-C(O)CH=CHCOOH (XI)}$  as an intermediate, and said method further comprises:

heating said  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-NH-C(O)CH=CHCOOH}$  under conditions effective to promote cyclization by elimination of water to form  $\text{POLY-[O]}_b\text{-C(O)-NH-X-MAL}$ .

95. (Withdrawn) The method of claim 92, further comprising recovering said  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-MAL}$  from the reaction mixture.

96. (Withdrawn) The method of claim 95, wherein said recovered  $\text{POLY-[O]}_b\text{-C(O)-H}_2\text{N-X-MAL}$  has a purity of greater than about 80 %.

97. (Withdrawn) The method of claim 81, wherein X is a saturated acyclic, cyclic or alicyclic hydrocarbon chain having a total of about 3 to about 20 carbon atoms.

98. (Withdrawn) The method of claim 81, wherein X is a linear saturated acyclic hydrocarbon chain.

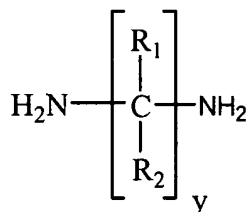
99. (Withdrawn) The method of claim 81, wherein X is a branched saturated acyclic hydrocarbon chain.

100. (Withdrawn) The method of claim 99, wherein X is branched at the carbon  $\alpha$  to the maleimidyl group.

101. (Withdrawn) The method of claim 99, wherein X is branched at the carbon  $\beta$  to the maleimidyl group.

102. (Withdrawn) The method of claim 99, wherein X is branched at the carbon  $\gamma$  to the maleimidyl group.

103. (Withdrawn) The method of claim 81, wherein said  $\text{NH}_2\text{-X-NH}_2$  corresponds to the structure:



**XII-A**

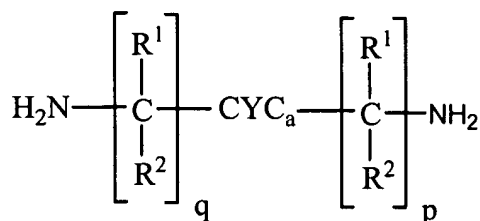
wherein

y is an integer from 1 to about 20;

$\text{R}^1$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl and

$\text{R}^2$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

104. (Withdrawn) The method of claim 81, wherein said  $\text{NH}_2\text{-X-NH}_2$  corresponds to the structure:



**XII-B**

and

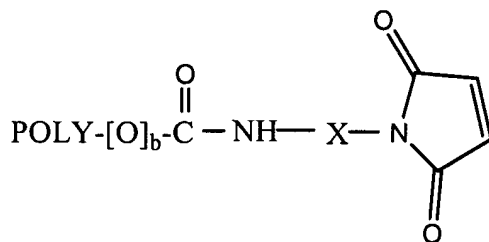
$\text{CYC}_a$  is a cycloalkylene group having "a" ring carbons, where the value of "a" ranges from 3 to 12;

p and q are each independently 0 to 20, and  $p + q + a \leq 20$ ,

$\text{R}^1$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and

$\text{R}^2$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

105. (Currently Amended) A conjugate formed by reaction of an active agent comprising ~~either a reactive thiol or amino-group~~ with a water-soluble polymer having the structure:



**II**

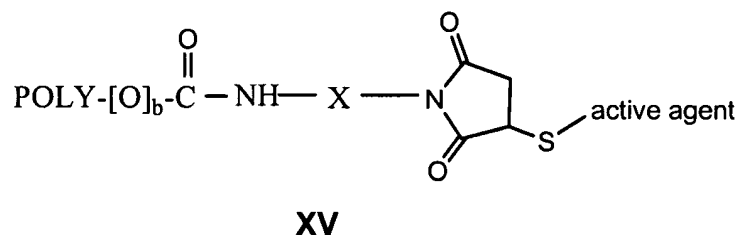
wherein:

POLY is a water-soluble polymer segment,

b is 0 or 1,

X is a hydrolytically stable linker comprising at least 4 contiguous saturated carbon atoms, and  
said polymer is absent aromatic groups and ester linkages.

106. (Previously Presented) A conjugate comprising the following structure:



wherein:

POLY is a water-soluble polymer segment,

b is 0 or 1,

X is a hydrolytically stable linker comprising at least 4 contiguous saturated carbon atoms,

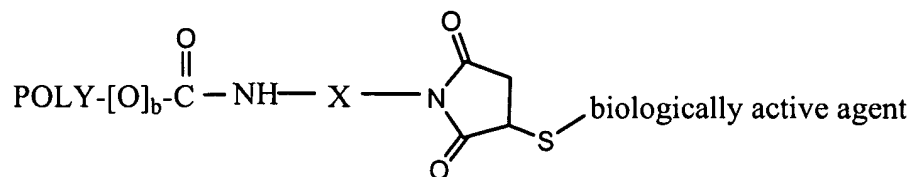
"POLY-[O]<sub>b</sub>-C(O)-NH-X-" is absent aromatic groups and ester linkages,  
and

"-S-active agent" represents a residue of an active agent comprising a thiol (-SH) group.

107. (Original) A composition comprising the conjugate of claim 106, wherein said composition comprises a single polymer conjugate species.

108. (Canceled)

109. (Withdrawn) A method for forming a polymer conjugate, said method comprising contacting a biologically active agent comprising a reactive thiol group, "HS-biologically active agent", with a polymer of claim 1, under conditions effective to promote formation of a polymer conjugate having the structure:



110. (Withdrawn) The method of claim 109, wherein said contacting step is carried out at pHs ranging from about 6.0 to about 8.0.

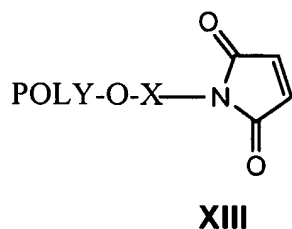
111. (Withdrawn) The method of claim 110, wherein said polymer conjugate is formed in a reaction mixture, and said method further comprises after said contacting step, isolating said polymer conjugate from said reaction mixture.

112. (Withdrawn) A hydrogel formed using the water-soluble polymer of claim 1.

113. (Withdrawn) A hydrogel formed using the water soluble polymer of claim 59.

114. (Withdrawn) A hydrogel formed using the water soluble polymer of claim 61.

115. (Withdrawn) A water soluble polymer having the structure:

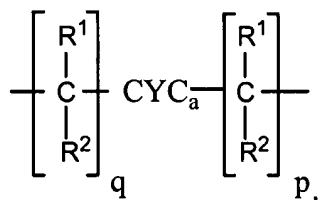


wherein:

POLY is a water-soluble polymer segment,

X is a hydrolytically stable linker that is a saturated cyclic or alicyclic hydrocarbon chain having a total of about 3 to about 20 carbon atoms,  
and  
said polymer is absent aromatic groups and ester linkages.

116. (Withdrawn) The polymer of claim 115, wherein X has the structure:



**XIII-A**

and

CYC<sub>a</sub> is a cycloalkylene group having "a" ring carbons, where the value of "a" ranges from 3 to 12;

p and q are each independently 0 to 20, and  $p + q + a \leq 20$ ,

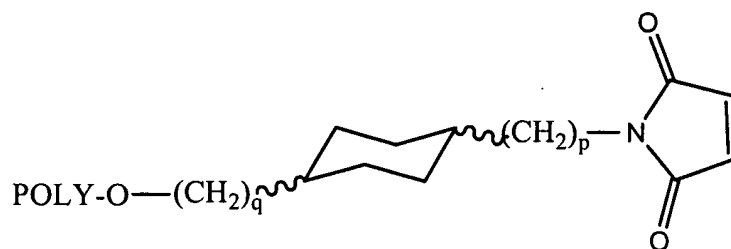
R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl,, and

R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl.

117. (Withdrawn) The polymer of claim 116, wherein p and q are each independently selected from the group consisting of 0, 1, 2, 3, 4, 5, 6, 7, and 8.

118. (Withdrawn) The polymer of claim 116, wherein R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of lower alkyl, lower cycloalkyl, and lower alkylenecycloalkyl, and R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of lower alkyl, lower cycloalkyl, and lower alkylenecycloalkyl.

119. (Withdrawn) The polymer of claim 116, wherein a is selected from the group consisting of 5, 6, 7, 8 and 9.
120. (Withdrawn) The polymer of claim 116, wherein the substituents on  $CYC_a$  are cis.
121. (Withdrawn) The polymer of claim 116, wherein the substituents on  $CYC_a$  are trans.
122. (Withdrawn) The polymer of claim 119, wherein a is 6 and  $CYC_a$  is a 1,1-, 1,2-, 1,3- or 1,4-substituted cyclohexyl ring.
123. (Withdrawn) The polymer of claim 116, wherein p and q each independently range from 0 to 4.
124. (Withdrawn) The polymer of claim 116, wherein  $R^1$  and  $R^2$  are H in every occurrence.
125. (Withdrawn) The polymer of claim 116, having the structure:

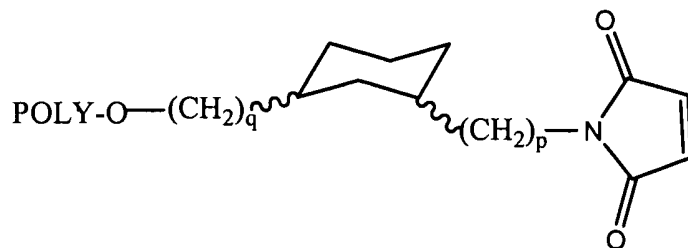
**XIII-B**

wherein q and p each independently range from 0 to 6.

126. (Withdrawn) The polymer of claim 116, wherein q ranges from 0 to 6 and p is zero.



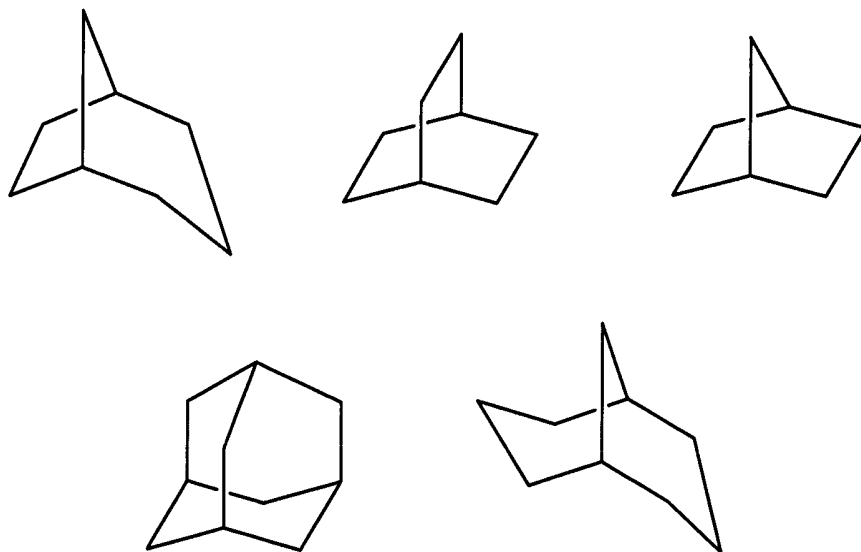
127. (Withdrawn) The polymer of claim 116, having the structure:



wherein q and p each independently range from 0 to 6.

128. (Withdrawn) The polymer of claim 116, wherein  $CYC_a$  is bicyclic or tricyclic.

129. (Withdrawn) The polymer of claim 116, wherein  $CYC_a$  is selected from the group consisting of:



and said ring substituents are positioned at any available position on the bi or tricyclic ring.

130. (Withdrawn) A method for forming a maleimide terminated polymer, said method comprising:

a. reacting  $\text{POLY-[O]}_b\text{-C(O)-LG}$  with  $\text{H}_2\text{N-X-MAL}$  under conditions effective to form  $\text{POLY-[O]}_b\text{-C(O)-HN-X-MAL}$ ,

wherein:

POLY is a water-soluble polymer segment,

b is 0 or 1,

X is a hydrolytically stable linker comprising at least 3 contiguous saturated carbon atoms,

LG is a leaving group,

MAL is maleimide,

and said maleimide terminated polymer is absent aromatic groups and ester linkages.

131. (Previously Presented) The conjugate of claim 105, wherein said active agent is selected from the group consisting of small molecules, peptides, and proteins.

132. (Previously Presented) The conjugate of claim 106, wherein said active agent is selected from the group consisting of small molecules, peptides, and proteins.

133. (Previously Presented) The conjugate of claim 106, wherein POLY is selected from the group consisting of poly(alkylene oxide), poly(vinyl pyrrolidone), poly(vinyl alcohol), polyoxazoline, poly(acryloylmorpholine), and poly(oxyethylated polyol).

134. (Previously Presented) The conjugate of claim 133, wherein POLY is a poly(alkylene oxide).

135. (Previously Presented) The conjugate of claim 134, wherein POLY is a poly(ethylene glycol).

136. (Previously Presented) The conjugate of claim 135, wherein the poly(ethylene glycol) is terminally capped with an end-capping moiety.

137. (Previously Presented) The conjugate of claim 136, wherein the end-capping moiety is selected from the group consisting alkoxy, substituted alkoxy, alkenyloxy, substituted alkenyloxy, alkynyloxy, substituted alkynyloxy, aryloxy, and substituted aryloxy.

138. (Previously Presented) The conjugate of claim 137, wherein the end-capping moiety is selected from the group consisting of methoxy, ethoxy, and benzyloxy.

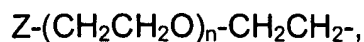
139. (Previously Presented) The conjugate of claim 135, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 100 daltons to about 100,000 daltons.

140. (Previously Presented) The conjugate of claim 139, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 1,000 daltons to about 50,000 daltons.

141. (Previously Presented) The conjugate of claim 140, wherein the poly(ethylene glycol) has a nominal average molecular mass of from about 2,000 daltons to about 30,000 daltons.

142. (Previously Presented) The conjugate of claim 135, wherein said poly(ethylene glycol) has a structure selected from the group consisting of linear, branched and forked.

143. (Previously Presented) The conjugate of claim 142, wherein said poly(ethylene glycol) comprises the structure:



where n is from about 10 to about 4000, and Z comprises a moiety selected from the group consisting of hydroxy, amino, ester, carbonate, aldehyde, aldehyde hydrate, acetal, ketone, ketone hydrate, ketal, alkenyl, acrylate, methacrylate, acrylamide, sulfone, thiol, carboxylic acid, isocyanate, isothiocyanate, hydrazide, urea, maleimide, vinylsulfone, dithiopyridine, vinylpyridine, iodoacetamide, alkoxy, benzyloxy, silane, lipid, phospholipid, biotin, and fluorescein.

144. (Previously Presented) The conjugate of claim 135, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, and 20.

145. (Previously Presented) The conjugate of claim 144, wherein X is a saturated acyclic, cyclic, or alicyclic hydrocarbon chain having a total number of carbon atoms selected from the group consisting of: from about 4 to about 12, from about 4 to about 10, and from about 5 to about 8 atoms.

146. (Previously Presented) The conjugate of claim 135, wherein X is a linear saturated acyclic hydrocarbon chain.

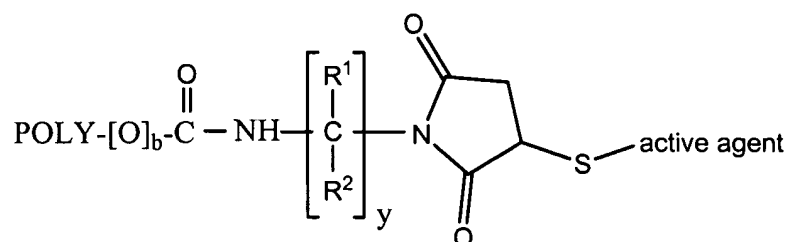
147. (Previously Presented) The conjugate of claim 135, wherein X is a branched saturated acyclic hydrocarbon chain.

148. (Previously Presented) The conjugate of claim 147, wherein X is branched at the carbon  $\alpha$  to the maleimidyl group.

149. (Previously Presented) The conjugate of claim 147, wherein X is branched at the carbon  $\beta$  to the maleimidyl group.

150. (Previously Presented) The conjugate of claim 147, wherein X is branched at the carbon  $\gamma$  to the maleimidyl group.

151. (Previously Presented) The conjugate of claim 146, having the structure:



wherein:

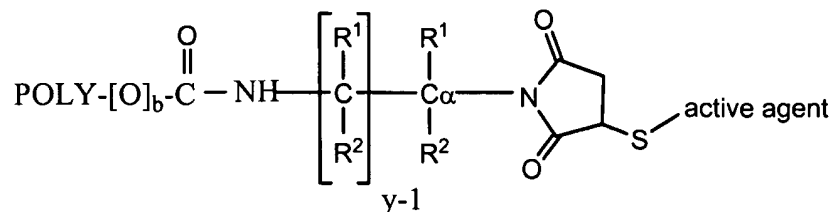
y is an integer from 4 to about 20;

$R^1$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl, and

$R^2$ , in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl.

152. (Previously Presented) The conjugate of claim 151, wherein either (i)  $R^1$  and  $R^2$  in each occurrence is independently H or an organic radical selected from the group consisting of lower alkyl and lower cycloalkyl or (ii)  $R^1$  and  $R^2$  are both H, and y is selected from the group consisting of 4, 5, 6, 7, 8, 9, and 10.

153. (Previously Presented) The conjugate of claim 151 having the structure:



wherein at least one of  $R^1$  or  $R^2$  on  $C_\alpha$  is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl.

154. (Previously Presented) The conjugate of claim 153, wherein each of  $R^1$  and  $R^2$  on  $C_\alpha$  is independently selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylenecycloalkyl, and substituted alkylenecycloalkyl.

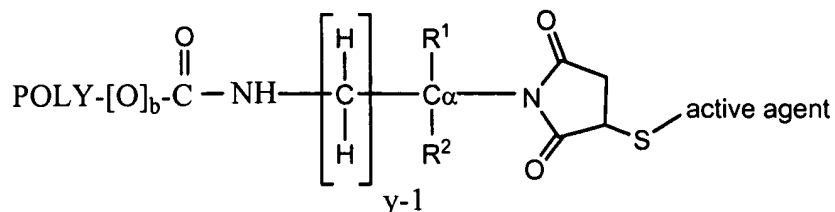
155. (Previously Presented) The conjugate of claim 153, wherein all other non- $C_\alpha$   $R^1$  and  $R^2$  variables are H.

156. (Previously Presented) The conjugate of claim 153 wherein at least one of  $R^1$  or  $R^2$  on  $C_\alpha$  is lower alkyl or lower cycloalkyl.

157. (Previously Presented) The conjugate of claim 153, wherein  $R^2$  on  $C_\alpha$  is H.

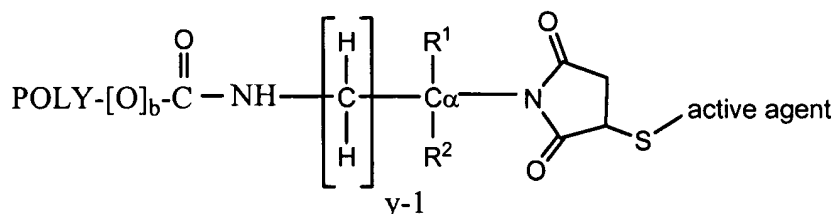
158. (Previously Presented) The conjugate of claim 157, wherein  $R^1$  on  $C_\alpha$  is selected from the group consisting of methyl, ethyl, propyl, isopropyl, butyl, isobutyl, pentyl, cyclopentyl, hexyl, and cyclohexyl.

159. (Previously Presented) The conjugate of claim 153 having the structure:



wherein  $\text{R}^1$  and  $\text{R}^2$  are each independently alkyl or cycloalkyl.

160. (Previously Presented) The conjugate of claim 153, having the structure:

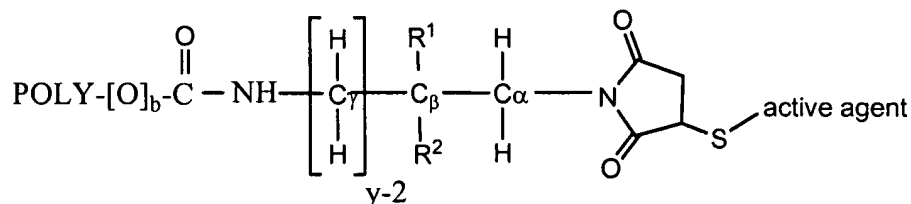


wherein  $\text{R}^1$  is alkyl or cycloalkyl and  $\text{R}^2$  is H.

161. (Previously Presented) The conjugate of claim 159 wherein  $\text{R}^1$  and  $\text{R}^2$  are each independently either methyl or ethyl.

162. (Previously Presented) The conjugate of claim 159, wherein  $\text{R}^1$  and  $\text{R}^2$  are the same.

163. (Previously Presented) The conjugate of claim 149 having the structure:



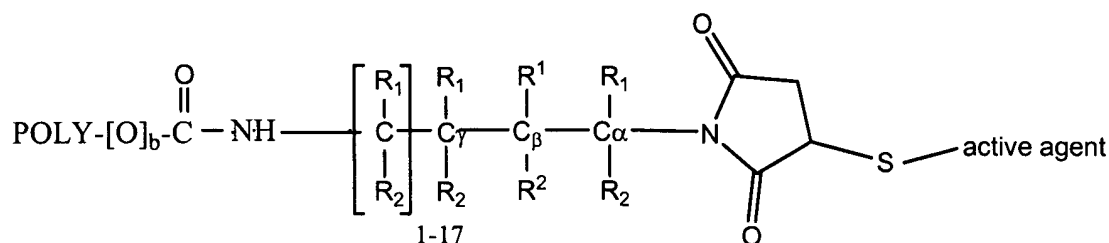
wherein  $R^1$  and  $R^2$  are each independently selected from the group consisting of H, alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkenecycloalkyl, and substituted alkenecycloalkyl, but are not both H.

164. (Previously Presented) The conjugate of claim 163, wherein  $R^1$  and  $R^2$  are each independently H, lower alkyl or lower cycloalkyl.

165. (Previously Presented) The conjugate of claim 164, wherein  $R^1$  and  $R^2$  are each independently selected from the group consisting of H, methyl, ethyl, propyl, isopropyl, butyl, isobutyl, pentyl, cyclopentyl, hexyl, and cyclohexyl.

166. (Previously Presented) The conjugate of claim 163, wherein  $R^2$  is H.

167. (Previously Presented) The conjugate of claim 151, having the structure:



wherein at least one of  $R^1$  and  $R^2$  attached to  $C_\gamma$  is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkenecycloalkyl, and substituted alkenecycloalkyl.

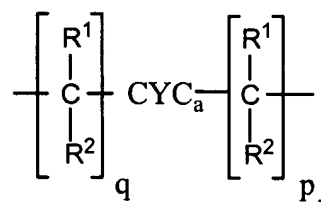
168. (Previously Presented) The conjugate of claim 167, wherein at least one of  $R^1$  and  $R^2$  attached to  $C_\gamma$  is alkyl or cycloalkyl and all other  $R^1$  and  $R^2$  variables are H.

169. (Previously Presented) The conjugate of claim 167, wherein one of the  $R^1$  variables attached to  $C_\alpha$  or  $C_\beta$  is alkyl or cycloalkyl, and all other  $R^1$  and  $R^2$  variables are H.



170. (Previously Presented) The conjugate of claim 106, wherein X is a saturated cyclic or alicyclic hydrocarbon chain.

171. (Previously Presented) The conjugate of claim 170, wherein X has the structure:



**IV**

and

CYC<sub>a</sub> is a cycloalkylene group having "a" ring carbons, where the value of "a" ranges from 3 to 12;

p and q are each independently 0 to 20, and  $p + q + a \leq 20$ ,

R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl, and

R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is selected from the group consisting of alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, alkylencycloalkyl, and substituted alkylencycloalkyl.

172. (Previously Presented) The conjugate of claim 171, wherein p and q are each independently selected from the group consisting of 0, 1, 2, 3, 4, 5, 6, 7, and 8.

173. (Previously Presented) The conjugate of claim 171, wherein R<sup>1</sup>, in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl, and R<sup>2</sup>, in each occurrence, is independently H or an organic radical that is either lower alkyl or lower cycloalkyl.

174. (Previously Presented) The conjugate of claim 171, wherein a is selected from the group consisting of 5, 6, 7, 8 and 9.

175. (Previously Presented) The conjugate of claim 174, wherein  $a$  is 6 and  $CYC_a$  is a 1,1-, 1,2-, 1,3- or 1,4-substituted cyclohexyl ring.

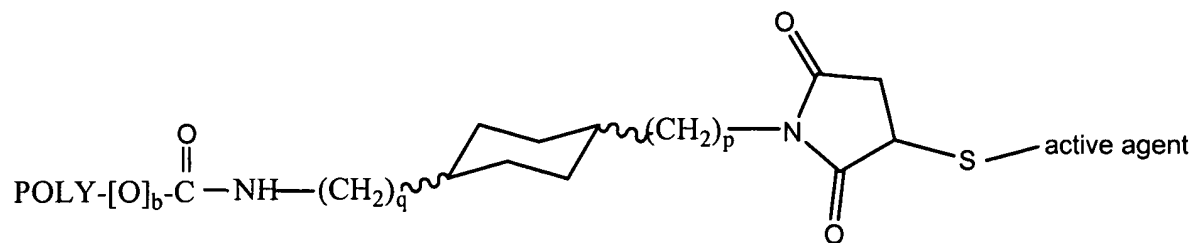
176. (Previously Presented) The conjugate of claim 171, wherein  $p$  and  $q$  each independently range from 0 to 4.

177. (Previously Presented) The conjugate of claim 175, wherein the substituents on said substituted cyclohexyl ring are *cis*.

178. (Previously Presented) The conjugate of claim 175, wherein the substituents on said substituted cyclohexyl ring are *trans*.

179. (Previously Presented) The conjugate of claim 171, wherein  $R^1$  and  $R^2$  are H in every occurrence.

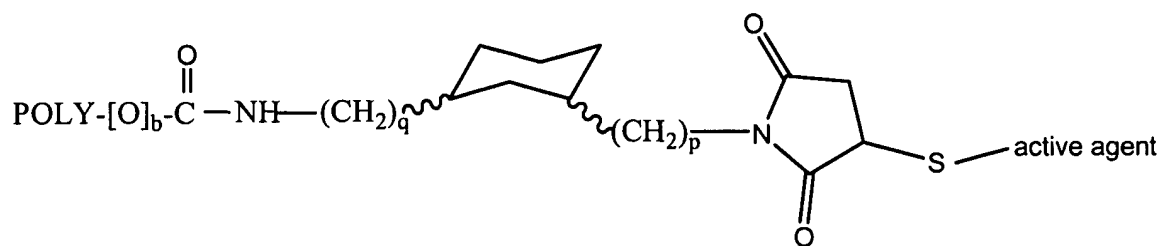
180. (Previously Presented) The conjugate of claim 173, having the structure:



wherein  $q$  and  $p$  each independently range from 0 to 6.

181. (Previously Presented) The conjugate of claim 180, wherein  $q$  ranges from 0 to 6 and  $p$  is zero.

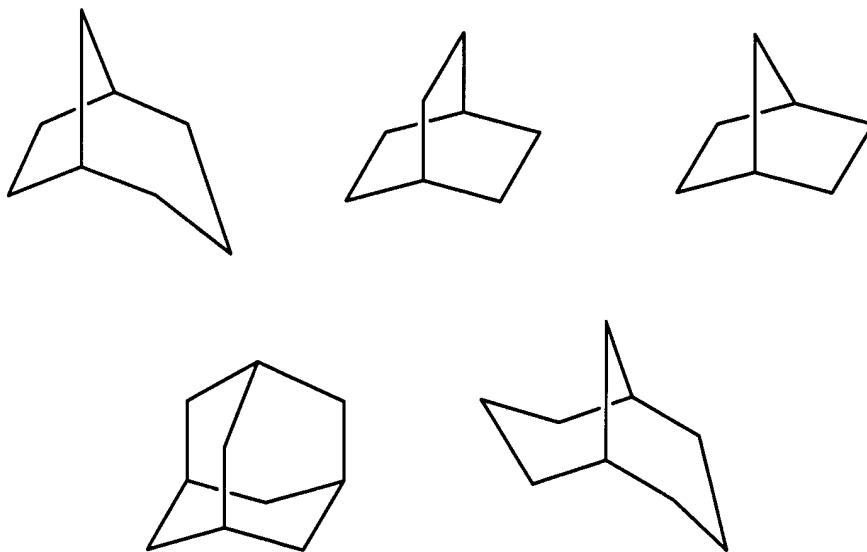
182. (Previously Presented) The conjugate of claim 173, having the structure:



wherein q and p each independently range from 0 to 3, and the substituents on the cyclohexylene ring are either cis or trans.

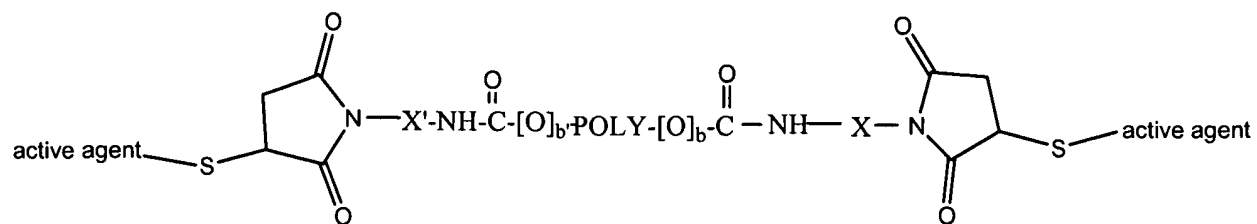
183. (Previously Presented) The conjugate of claim 171, wherein CYC<sub>a</sub> is bicyclic or tricyclic.

184. (Previously Presented) The conjugate of claim 183, wherein CYC<sub>a</sub> is selected from the group consisting of:



and said ring substituents are positioned at any available position on the bi or tricyclic ring.

185. (Previously Presented) The conjugate of claim 106 having the structure:

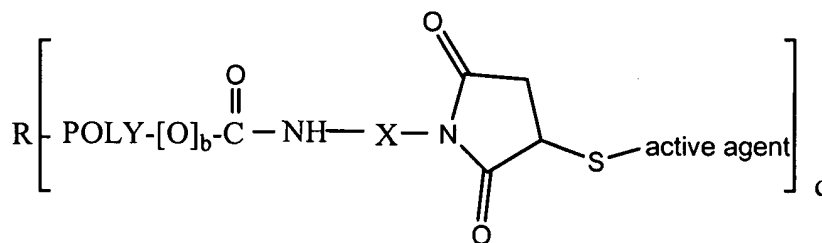


wherein X and b are as previously defined,

b' is 0 or 1, and

X' is a hydrolytically stable linker comprising at least 4 contiguous saturated carbon atoms.

186. (Previously Presented) The conjugate of claim 106, corresponding to the structure:



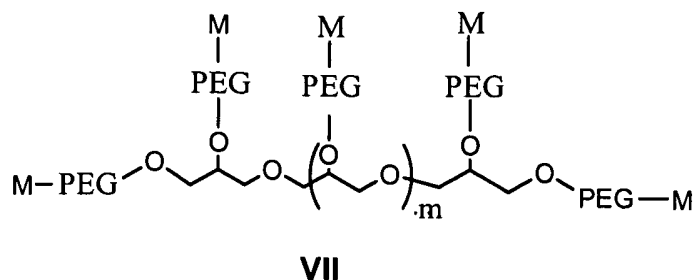
wherein:

d is an integer from 3 to about 100, and

R is a residue of a central core molecule having 3 or more hydroxyl groups, amino groups, or combinations thereof.

187. (Previously Presented) The conjugate of claim 186, wherein d is an integer from 3 to about 12.

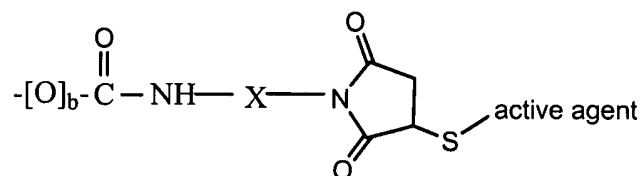
188. (Previously Presented) The conjugate of claim 105, wherein POLY is a multi-arm polymer segment, and said polymer corresponds to the structure:



where

PEG is  $-(\text{CH}_2\text{CH}_2\text{O})_n\text{CH}_2\text{CH}_2-$ ,

M is :



and m is selected from the group consisting of 3, 4, 5, 6, 7, and 8.